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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/910,383	07/20/2001	Girish N. Nallur	13172.0007U1	2312

23859 7590 12/13/2006

NEEDLE & ROSENBERG, P.C.  
SUITE 1000  
999 PEACHTREE STREET  
ATLANTA, GA 30309-3915

EXAMINER

CALAMITA, HEATHER

ART UNIT	PAPER NUMBER
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1637

DATE MAILED: 12/13/2006

Please find below and/or attached an Office communication concerning this application or proceeding.

<b>Office Action Summary</b>	<b>Application No.</b>	<b>Applicant(s)</b>	
	09/910,383	NALLUR ET AL.	
	<b>Examiner</b>	<b>Art Unit</b>	
	Heather G. Calamita, Ph.D.	1637	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --  
**Period for Reply**

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

#### Status

- 1) ☒ Responsive to communication(s) filed on 02 October 2006.
- 2a) ☒ This action is **FINAL**.                      2b) ☐ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

#### Disposition of Claims

- 4) ☒ Claim(s) 1-62, 68-76 and 82-87 is/are pending in the application.
- 4a) Of the above claim(s) \_\_\_\_\_ is/are withdrawn from consideration.
- 5) ☐ Claim(s) \_\_\_\_\_ is/are allowed.
- 6) ☒ Claim(s) 1-62, 68-76 and 82-87 is/are rejected.
- 7) ☐ Claim(s) \_\_\_\_\_ is/are objected to.
- 8) ☐ Claim(s) \_\_\_\_\_ are subject to restriction and/or election requirement.

#### Application Papers

- 9) ☐ The specification is objected to by the Examiner.
- 10) ☐ The drawing(s) filed on \_\_\_\_\_ is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.  
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).  
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
- 11) ☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

#### Priority under 35 U.S.C. § 119

- 12) ☐ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
- a) ☐ All    b) ☐ Some \* c) ☐ None of:
1. ☐ Certified copies of the priority documents have been received.
  2. ☐ Certified copies of the priority documents have been received in Application No. \_\_\_\_\_.
  3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).
- \* See the attached detailed Office action for a list of the certified copies not received.

#### Attachment(s)

- |  |   |
|--|---|
| 1) <input type="checkbox"/> Notice of References Cited (PTO-892)                     | 4) <input type="checkbox"/> Interview Summary (PTO-413)           |
| 2) <input type="checkbox"/> Notice of Draftsperson's Patent Drawing Review (PTO-948) | Paper No(s)/Mail Date. _____                                      |
| 3) <input type="checkbox"/> Information Disclosure Statement(s) (PTO/SB/08)          | 5) <input type="checkbox"/> Notice of Informal Patent Application |
| Paper No(s)/Mail Date _____  | 6) <input type="checkbox"/> Other: _____                          |

## DETAILED ACTION

### *Status of Application, Amendments, and/or Claims*

1. Claims 1-62, 68-76 and 82-87 are pending and under examination. All arguments have been fully considered and thoroughly reviewed, but are deemed not persuasive for the reasons that follow. Any objections and rejections not reiterated below are hereby withdrawn.

### *Claim Rejections - 35 USC § 103*

2. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

(a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negated by the manner in which the invention was made.

Claims 1-15, 18-29, 31-47, 53-58, 61, 62, 68, 70-73 and 82-87 are rejected under 35 U.S.C. 103(a) as being unpatentable over Lizardi et al. (USPN 6,316,229 B1) in view of Schweitzer et al. (PNAS 2000).

Lizardi et al. teach (claims 1, 53, 56, 62, 68, 70-73 and 87) a method of amplifying messenger RNA, the method comprising (see entire document, specifically col. 42 line 13).

(a) mixing one or more RT primers with a nucleic acid sample and reverse transcribing to produce cDNA strands each comprising one of the RT primers, wherein each RT primer comprises a reverse transcription primer portion (see col. 77 line 2, where Lizardi et al. indicate RNA can be used with the methodologies disclosed).

(b) mixing the cDNA strands with a set of capture probes under conditions that promote hybridization of the cDNA strands to the capture probes (see col. 42 lines 27-52)

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(c) mixing one or more rolling circle replication primers with the cDNA strands under conditions that promote association of the cDNA strands with the rolling circle replication primers, wherein the rolling circle replication primers (see col. 42 lines 27-52),

(d) mixing one or more amplification target circles with the rolling circle replication primers under conditions that promote association of the rolling circle replication primers with the amplification target circles (see col. 42 lines 53-67),

(e) incubating the amplification target circles under conditions that promote replication of the amplification target circles (see col. 42 lines 65-67, col. 43 lines 1-4), wherein replication of the amplification target circles results in the formation of tandem sequence DNA (see col. 43 lines 5-20).

With regard to claim 3, Lizardi et al. teach the reverse transcription primer portion of each RT primer comprises poly T (see col. 77 line 65).

With regard to claim 4, Lizardi et al. teach the capture probes are immobilized on a substrate (see col. 42 lines 29-30).

With regard to claim 5, Lizardi et al. teach the capture probes are in an array (see col. 42 line 8).

With regard to claim 6, Lizardi et al. teach the capture probes are immobilized via a capture tag coupled to the capture probes (see example 5 col. 78 steps 1 and 2).

With regard to claim 7, Lizardi et al. teach each capture probe comprises a sequence matching all or a portion of the sequence of messenger RNA molecules of interest (see col. 77 line 65, col. 78 line 4).

With regard to claims 8-12, Lizardi et al. teach the set of capture probes collectively comprise sequence matching all or a portion of the sequence of a plurality of different messenger RNA molecules associated with a disease from a source of interest (see col. 51 lines 63-67, col. 52 lines 1-40).

With regard to claim 13, Lizardi et al. teach the ends of the capture probes are extendable when a cDNA strand is hybridized to the capture probe (see col. 42 line 67).

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With regard to claim 14, Lizardi et al. teach the ends of the capture probes are designed to be extendable only when a cDNA strand corresponding to a particular form of a messenger RNA of interest is hybridized to the capture probe (see col. 42 line 67).

With regard to claim 15, Lizardi et al. teach the ends of the capture probes are not extendable by polymerase (see col. 47 lines 62-64).

With regard to claim 18, Lizardi et al. teach further comprising, simultaneous with, or following, step (d), mixing a secondary DNA strand displacement primer with the amplification target circles and incubating under conditions that promote hybridization between the tandem sequence DNA and the secondary DNA strand displacement primer and replication of the tandem sequence DNA, wherein replication of the tandem sequence DNA results in the formation of secondary tandem sequence DNA (see col. 55 lines 40-54).

With regard to claim 19, Lizardi et al. teach further comprising, simultaneous with step (e), mixing a tertiary DNA strand displacement primer with the amplification target circles (see col. 55 lines 1-10).

With regard to claim 20, Lizardi et al. teach further comprising detecting the tandem sequence DNA, wherein detection of tandem sequence DNA indicates that the corresponding messenger RNA molecule was present in the nucleic acid sample (see col. 52 lines 5-27).

With regard to claim 21, Lizardi et al. teach the tandem sequence DNA is detected while in association with the capture probes.

With regard to claim 22, Lizardi et al. teach the identity of the capture probe associated with a tandem sequence DNA indicates the identity of the corresponding messenger RNA molecule (see col. 52 lines 5-27).

With regard to claim 23, Lizardi et al. teach the tandem sequence DNA is detected at the site where the capture probe is located, and wherein the location of the capture probe indicates the identity of the corresponding messenger RNA molecule (see col. 52 lines 5-27).

With regard to claim 24, Lizardi et al. teach detection is mediated by detection probes or by a detection label incorporated in the tandem sequence DNA (see col. 48 lines 48-54).

With regard to claim 25, Lizardi et al. teach the detection label is a ligand (see col. 48 line 47).  
With regard to claim 26, Lizardi et al. teach the ligand is Brdu (see col. 48 lines 32-33).

With regard to claim 27, Lizardi et al. teach the ligand is Brdu, wherein the tandem sequence DNA is detected by associating an anti-Brdu antibody with the tandem sequence DNA and detecting the antiBrdu antibody (see col. 48 line 33).

With regard to claim 28, Lizardi et al. teach the anti-Brdu antibody comprises a label, wherein the anti-Brdu antibody is detected by detecting the label (see col. 48 lines 32-33).

With regard to claim 29, Lizardi et al. teach the label on the anti-Brdu antibody is a fluorophore (see col. 48 lines 32-37).

With regard to claim 31, Lizardi et al. teach further comprising mixing a set of detection probes with the tandem sequence DNA under conditions that promote hybridization between the tandem sequence DNA and the detection probes, and detecting a plurality of different sequences present in the tandem sequence DNA (see col. 61 lines 9-21).

With regard to claim 32, Lizardi et al. teach the tandem sequence DNA is collapsed using collapsing probes (see col. 52 lines 60-67).

With regard to claim 33, Lizardi et al. teach at least one of the collapsing probes is a collapsing detection probe (see col. 52 lines 22-23, 60).

With regard to claim 34, Lizardi et al. teach the tandem sequence DNA is collapsed by mixing the collapsing probes with the tandem sequence DNA, and incubating under conditions that promote

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hybridization between the collapsing probes and the tandem sequence DNA (see col. 52 lines 66-67, col. 55 line 14).

With regard to claim 35, Lizardi et al. teach further comprising, prior to or simultaneous with the mixing of the collapsing probes with the tandem sequence DNA, mixing detection probes with the tandem sequence DNA, and incubating under conditions that promote hybridization between the detection probes and the tandem sequence DNA (see col. 62 lines 66-67, col. 63 lines 1-10).

With regard to claim 36, Lizardi et al. teach the collapsing probes comprise ligands, haptens, or both coupled to or incorporated into oligonucleotides (see col. 63 lines 50-53, col. 24 lines 65-67).

With regard to claim 37, Lizardi et al. teach the RT primer comprises a capture tag (see col. 23 lines 50-67).

With regard to claims 38, 41 and 46, Lizardi et al. teach the biotin (see col. 23 lines 50-67).

With regard to claims 43 and 44, Lizardi et al. teach the association occurs between a protein and a nucleic acid (see col. 78 lines 31-32).

With regard to claim 61, Lizardi et al. teach the incorporation of biotinylated-ddNTP into the cDNA (see col. 43 lines 10-20).

Lizardi et al. do not teach the capture tag is a hapten a ligand a ligand binding molecule an antibody or an anti-antibody and the capture tag is not a nucleic acid.

Schweitzer et al. teach the capture tag is an antibody.

With regard to claim 2, Schweitzer et al. teach the capture tag associates with the primer (see the abstract, where the primer is covalently attached to the antibody).

With regard to claims 39 and 40, Schweitzer et al. teach the DNA strands comprise capture tags (see the abstract, where the primer is covalently attached to the antibody).

With regard to claim 42, Schweitzer et al. teach the association is covalent (see the abstract, where the primer is covalently attached to the antibody).

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With regard to claim 45, Schweitzer et al. teach the association occurs between two proteins (see p. 10116 col. 2 2<sup>nd</sup> full paragraph, where the analyte is PSA, a protein, and the tag bound to the primer is anti-PSA antibody, also a protein).

With regard to claims 54 and 57, Schweitzer et al. teach the rolling circle replication primers each comprise a capture tag (see the abstract, where the primer is covalently attached to the antibody).

With regard to claims 55 and 58, Schweitzer et al. teach association of the rolling circle replication primers with the DNA occurs via association of the capture tag added to the DNA and the capture tag in the rolling circle replication primers (see the abstract, where the primer is covalently attached to the antibody).

One of ordinary skill in the art at the time the invention was made would have been motivated to use the method of attaching an antibody capture tag to a primer, as taught by Schweitzer with the method of amplifying target nucleic acids as taught by Lizardi in order to have a versatile ultrasensitive method of antigen detection. Schweitzer teaches that the RCA reporter system can be adapted for the detection of protein antigens using an oligonucleotide primer that is covalently attached to an antibody. Schweitzer additionally, teaches using a single primer, RCA generates hundreds of tandemly linked copies of the circular template within a few minutes and the 5' end of the primer is attached to an antibody. The amplified DNA can be detected in a variety of ways including direct incorporation of hapten labeled or fluorescently labeled nucleotides. An ordinary practitioner would have been motivated to use the method of attaching an antibody capture tag to a primer, as taught by Schweitzer with the method of amplifying target nucleic acids as taught by Lizardi in order to expand the genus of analytes which can be detected using RCA.

3. Claims 16 and 17 are rejected under 35 U.S.C. 103(a) as being unpatentable over Lizardi et al. (USPN 6,316,229 B1) and Schweitzer et al. (PNAS 2000) in view Lizardi (US 2003/0032024 A1).



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The teachings and suggestions of Lizardi (229) and Schweitzer are described previously.

Lizardi (229) and Schweitzer do not teach or suggest subprobes.

With regard to claims 16 and 17, Lizardi (024) teaches mixing one or more subprobes (gap oligonucleotides) with the cDNA strands wherein each half probe is designed to hybridize to a cDNA strand adjacent to where a capture probe hybridizes, ligating the subprobes and capture probes hybridized, and after ligation, incubating the capture probes at a temperature above the melting temperature of the capture probe but below the melting temperature of the ligated capture probe/subprobe (see paragraph 0195).

One of ordinary skill in the art at the time the invention was made one would have been motivated to apply Gap oligonucleotides as taught by Lizardi (024) with the method of amplifying target nucleic acids as taught by Lizardi (229) and Schweitzer in order to achieve more selective target discrimination. Lizardi (024) teaches using gap oligonucleotides enhance target dependency in LCR and this can be adapted for use in LM-RCA, and this method enhances target discrimination. It would have been prima facie obvious to use the gap oligonucleotide as taught by Lizardi (024) with the method of amplifying nucleic acids as taught by Lizardi (229) and Schweitzer in order to achieve the expected advantage of enhancing target nucleic acid discrimination.

4. Claim 30 is rejected under 35 U.S.C. 103(a) as being unpatentable over Lizardi et al. (USPN 6,316,229 B1) and Schweitzer et al. (PNAS 2000) in view of Waggoner et al. (USPN 6,008,373).

The teachings and suggestions of Lizardi (229) Schweitzer are described previously.

Lizardi (229) and Schweitzer do not teach or suggest phycoerythrin as a fluorophore.

Waggoner et al. teach using phycoerythrin as a fluorophore in the detection label on an antibody (see col. 21 line 64).

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One of ordinary skill in the art at the time the invention was made one would have been motivated to use phycoerythrin as taught by Waggoner with the method of amplifying target nucleic acids as taught by Lizardi (229) and Schweitzer to achieve a detection signal that provides fluorescence that is relatively free of interference from other biological materials and provides a multicolor fluorescence emission using a single wavelength excitation. Waggoner teach phycoerythrin is advantageous because it is low molecular weight and provides a multicolor fluorescence emission using a single wavelength excitation (see col. 2 lines 28-30). It would have been prima facie obvious to use Phycoerythrin as taught by Waggoner with the method of amplifying nucleic acids as taught by Lizardi (229) and Schweitzer in order to achieve the expected advantage of a label that has a low molecular weight and provides a multicolor fluorescence emission from a single excitation wavelength.

5. Claims 48-52, and 69 are rejected under 35 U.S.C. 103(a) as being unpatentable over Lizardi et al. (USPN 6,316,229 B1) and Schweitzer et al. (PNAS 2000) in view of Cao et al. (US 2002/0120409 A1).

The teachings and suggestions of Lizardi (229) Schweitzer are described previously.

Lizardi (229) and Schweitzer do not teach or suggest fragmenting and labeling cDNA strands to form labeled fragmented cDNA.

Cao et al. teach fragmented cDNA in a method to amplify mRNA (see claim 1 page 8).

One of ordinary skill in the art at the time the invention was made one would have been motivated to use the method of fragmenting and labeling cDNA as taught by Cao et al. with the method of amplifying target nucleic acids as taught by Lizardi (229) and Schweitzer to obtain labeled cDNA fragments that are used in assessing gene expression. It would have been prima facie obvious to use fragmenting and labeling cDNA as taught by Cao et al. with the method of amplifying target nucleic acids

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as taught by Lizardi (229) and Schweitzer in order to achieve the expected advantage of using the labeled cDNA fragments in gene expression arrays.

6. Claims 59-60 are rejected under 35 U.S.C. 103(a) as being unpatentable over Lizardi et al. (USPN 6,316,229 B1) and Schweitzer et al. (PNAS 2000) in view of Shoemaker et al. (USPN 6,713,257 B2).

The teachings and suggestions of Lizardi (229) and Schweitzer are described previously.

Lizardi (229) and Schweitzer do not teach or suggest teach a capture tag derived from allyl amine dUTP.

Shoemaker et al. teaches using an amino-allyl dUTP in labeling cDNA (see col. 34 line 8).

One of ordinary skill in the art at the time the invention was made one would have been motivated to use the method of labeling cDNA as taught by Shoemaker et al. with the method of amplifying target nucleic acids as taught by Lizardi (229) and Schweitzer to obtain labeled cDNA that are used in assessing gene expression. It would have been prima facie obvious to use the method of labeling cDNA as taught by Shoemaker et al. with the method of amplifying target nucleic acids as taught by Lizardi (229) and Schweitzer in order to achieve the expected advantage of incorporating a detectible fluorescent label into the cDNA of interest.

### ***Response to Arguments***

7. Applicants' arguments filed October 2, 2006, have been fully considered but they are not persuasive.

Applicants' arguments with respect to claims 1-15, 18-29, 53-58, 61, 62, 68, 70-73 and 82-87 are not persuasive. Applicants argue the claims require either the rolling circle replication primer comprises a capture tag or the cDNA comprises a capture tag and the capture tag is a hapten, ligand, ligand binding

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molecule, an antibody or an anti-antibody and that the 229 patent does not teach this. This is not persuasive because the 229 patent was not relied on for this teaching. Schweitzer et al. was relied on for the teaching of a capture tag that is an antibody. Schweitzer et al. teach this in see the abstract, where the primer is covalently attached to the antibody as stated above, specifically a nucleic acid tag covalently coupled to an antibody. Applicants' further argue the claims require the association between the rolling circle replication primers and the cDNA occurs via the capture tag and the 229 patent teaches an association of the primer and the target DNA molecule occurs via a nucleotide to nucleotide base pairing interaction between the sequences and not by interaction of a hapten or ligand. This is not persuasive because the nucleotide to nucleotide interaction meets the generic limitation of capture tag, as a nucleotide can be a capture tag. Further the specific requirement in the claim defining the capture tag recites "capture tag is a hapten, ligand, ligand binding molecule, an antibody or an anti-antibody" the 229 patent was not relied on to meet this specific limitation rather Schweitzer et al. which teaches a capture tag of an antibody was relied on. Applicants further argue no motivation for combining the references. This is not persuasive because it would be obvious to substitute an antibody for the nucleotide tag because it would improve specificity and enhance target discrimination. Additionally the 229 patent expressly teaches a hapten a ligand and a ligand binding molecule at col. 53 lines 54-56. Applicants further argue the combination of the 229 patent and Schweitzer would render the method of the 229 patent inoperable. This is not persuasive. The combination of Schweitzer with the 229 patent would simply change the operation of the methods disclosed in the 229 patent. One of skill in the art recognizes there exists a variety of ways to capture in a specific manner and the 024 patent provides motivation indicating that gap oligonucleotides enhance target dependency in LCR and this can be adapted for use in LM-RCA, and this method enhances target discrimination. Finally, Applicants argue Schweitzer do not teach the use of RT primers, cDNA formation or amplifying messenger RNA. This is not persuasive because Schweitzer was

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not relied on for these teachings. The 229 patent discloses all of these claim elements and therefore was relied on for these teachings.

Applicants' arguments with respect to claims 37-41, 53-55, 62, 70 and 72 are not persuasive. Applicants argue the 229 patent fails to disclose or refer specifically to RT primers. This is not persuasive because patent 229 expressly states that RNA can be used with the methodologies disclosed. Therefore a practitioner of ordinary skill in the art recognizes the need for using RT primers when working with RNA. Additionally the 229 patent teaches cDNA which inherently requires the use of RT primers. Moreover primers are primers. RT primers (primers for producing cDNA) are not structurally different from DNA primers (primers for producing DNA), therefore because the combination of the 229 patent and 024 patent renders the instant invention obvious with respect to DNA primers then it necessarily renders the instant invention obvious with respect to RT primers.

Applicants' additional arguments for claims 39-41 are not persuasive. Applicants argue the 229 patent fails to teach the cDNA strands comprise a capture tag and the capture tag is selected from the group consisting of biotin, digoxigenin, bromodeoxyuridine or other haptens. Applicants argue while the 229 patent discloses the capture tags, the tags are not disclosed on cDNA. This is not persuasive because the 229 patent discloses RNA and further discloses cDNA (see example 4).

With respect to claims 46 and 47, Applicants note the office action does not address claim 47. This is an inaccurate characterization as the claim 47 discloses the limitation of "antibodies which bind biotin" the limitation of biotin is addressed at p. 6 of the office action and the limitation of the use of antibodies is also addressed at p. 6, therefore while "claim 47" is not specifically recited the limitations of claim 47 are addressed. With respect to Applicants' arguments regarding these claims they are not persuasive for reasons outlined above.

With respect to claims 83-85, Applicants assert the limitations of these claims were not addressed in the Office Action. Again this is an inaccurate characterization as claim 83 recites the limitation of

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“wherein the capture tag is a hapten, a ligand, a ligand binding molecule, an antibody, or an anti-antibody” and claim 84 recites the limitation of “wherein the capture tag is a hapten, an antibody or an anti-antibody” and claim 85 recites the limitation of “wherein the capture tag is not a nucleic acid” these limitations were addressed at p.6 of the Office action where the action states the 229 patent does not teach a capture tag of an antibody and that Schweitzer et al. teach this limitation. Additionally, Schweitzer teach an antibody as a capture therefore Schweitzer teaches the capture tag is not a nucleic acid as an antibody is not a nucleic acid. With respect to Applicants’ arguments regarding these claims they are not persuasive for reasons outlined above.

Applicants' arguments with respect to claim 30 are not persuasive. Applicants reiterate previously stated arguments relating to the deficiencies of the combination of the 229 and Schweitzer and additionally state that Waggoner et al. fail to supplement the elements missing from the 229 and 024 patents. This is not persuasive for the reasons outlined above.

Applicants' arguments with respect to claims 48-52, 69 and 73 are not persuasive. Applicants reiterate previously stated arguments relating to the deficiencies of the combination of the 229 and Schweitzer patents and additionally state that Cao et al. fail to supplement the elements missing from the 229 and 024 patents. This is not persuasive for the reasons outlined above.

Applicants' arguments with respect to claim 59 and 60 are not persuasive. Applicants reiterate previously stated arguments relating to the deficiencies of the combination of the 229 patent and Schweitzer and additionally state that Shoemaker et al. fail to supplement the elements missing from the 229 patent and Schweitzer. This is not persuasive for the reasons outlined above.

### ***Conclusion***

8. **THIS ACTION IS MADE FINAL.** Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

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A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the mailing date of this final action.

#### *Correspondence*

9. Any inquiry concerning this communication or earlier communications from the examiner should be directed to Heather G. Calamita whose telephone number is 571.272.2876 and whose e-mail address is heather.calamita@uspto.gov. However, the office cannot guarantee security through the e-mail system nor should official papers be transmitted through this route. The examiner can normally be reached on Monday through Thursday, 7:00 AM to 5:30 PM.

If attempts to reach the examiner are unsuccessful, the examiner's supervisor, Gary Benzion can be reached at 571.272.0782.

Papers related to this application may be faxed to Group 1637 via the PTO Fax Center using the fax number 571.273.8300.

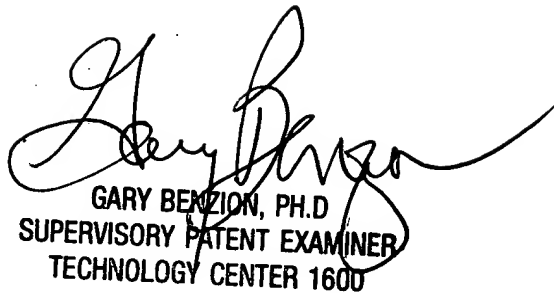
Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to 571.272.0547.

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GARY BENZION, PH.D  
SUPERVISORY PATENT EXAMINER  
TECHNOLOGY CENTER 1600